

**TRANSMITTAL LETTER TO THE UNITED STATES  
DESIGNATED/ELECTED OFFICE (DO/EO/US)  
CONCERNING A FILING UNDER 35 U.S.C. 371**

U.S. APPLICATION NO. (If known, see 37 CFR 1.5)

**10/501 176**

|                               |                           |                       |
|-------------------------------|---------------------------|-----------------------|
| INTERNATIONAL APPLICATION NO. | INTERNATIONAL FILING DATE | PRIORITY DATE CLAIMED |
| PCT/US03/01120                | 1/14/2003                 | 1/18/2002             |

## TITLE OF INVENTION

SELECTIVE S1P1/EDG1 RECEPTOR AGONISTS

## APPLICANT(S) FOR DO/BO/US

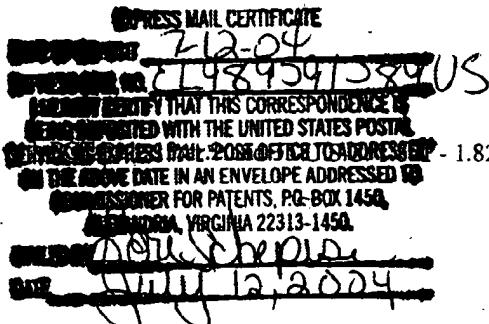
George A. Doherty, Michael J. Forrest, Richard Hajdu, Li Zhen, Jeffrey J. Hale, Suzanne M. Mandala, Sander G. Mills, Hugh Rosen and Edward M. Scolnick

**Applicant herewith submits to the United States Designated/Elected Office (DO/EO/US) the following items and other information:**

1.  This is a **FIRST** submission of items concerning a filing under 35 U.S.C. 371.
2.  This is a **SECOND** or **SUBSEQUENT** submission of items concerning a filing under 35 U.S.C. 371.
3.  This is an express request to begin national examination procedures [35 U.S.C. 371(f)]. The submission must include items (5), (6), (9) and (19) indicated below.
4.  The US has been elected (PCT Article 31).
5.  A copy of the International Application as filed [35 U.S.C. 371(c)(2)]
  - a.  is attached hereto (required only if not communicated by the International Bureau).
  - b.  has been communicated by the International Bureau.
  - c.  is not required, as the application was filed in the United States Receiving Office (RO/US).
6.  An English language translation of the International Application as filed [35 U.S.C. 371(c)(2)].
7.  Amendments to the claims of the International Application under PCT Article 19 [35 U.S.C. 371(c)(3)].
  - a.  are attached hereto (required only if not communicated by the International Bureau).
  - b.  have been communicated by the International Bureau.
  - c.  have not been made; however, the time limit for making such amendments has NOT expired.
  - d.  have not been made and will not be made.
8.  An English language translation of the amendments to the claims under PCT Article 19 [35 U.S.C. 371(c)(3)].
9.  An oath or declaration of the inventor(s) [35 U.S.C. 371(c)(4)].
10.  An English language translation of the annexes to the International Preliminary Examination Report under PCT Article 36 [35 U.S.C. 371(c)(5)].

**Items 11 to 18 below concern other document(s) or information included:**

11.  An Information Disclosure Statement under 37 CFR 1.97 and 1.98.
12.  An assignment document for recording. A separate cover sheet in compliance with 37 CFR 3.28 and 3.31 is included.
13.  A **FIRST** preliminary amendment.
14.  A **SECOND** or **SUBSEQUENT** preliminary amendment.
15.  A substitute specification.
16.  A power of attorney and/or change of address letter.
17.  A computer-readable form of the sequence listing in accordance with 37 CFR 1.825.
18.  Other items or information:



|   |   |   |                     |               |                  |                 |
|---|---|---|---------------------|---------------|------------------|-----------------|
| U.S. APPLICATION NO. (If known, see 37 CFR 1.5)<br><b>10/501 176</b>  | INTERNATIONAL APPLICATION NO.<br>PCT/US03/01120 | ATTORNEY'S DOCKET NUMBER<br>21014YP   |                     |               |                  |                 |
| <p>19. <input checked="" type="checkbox"/> The following fees are submitted:</p> <p><b>BASIC NATIONAL FEE [37 CFR 1.492(a)(1)-(5)]:</b></p> <p>Neither international preliminary examination fee (37 CFR 1.482) nor international search fee [37 CFR 1.445(a)(2)] paid to USPTO and International Search Report not prepared by the EPO or JPO..... <b>\$1,080.00</b></p> <p>International preliminary examination fee (37 CFR 1.482) not paid to USPTO but International Search Report prepared by the EPO or JPO..... <b>\$920.00</b></p> <p>International preliminary examination fee (37 CFR 1.482) not paid to USPTO but international search fee [37 CFR 1.445(a)(2)] paid to USPTO ..... <b>\$770.00</b></p> <p>International preliminary examination fee paid to USPTO (37 CFR 1.482) but all claims did not satisfy provisions of PCT Article 33(1)-(4)..... <b>\$730.00</b></p> <p>International preliminary examination fee paid to USPTO (37 CFR 1.482) and all claims satisfied provisions of PCT Article 33(1)-(4)..... <b>\$100.00</b></p> |   | <b>CALCULATIONS</b>   | <b>PTO USE ONLY</b> |               |                  |                 |
|   |   | <b>ENTER APPROPRIATE BASIC FEE AMOUNT</b> = <b>\$100.00</b>   |                     |               |                  |                 |
|   |   | Surcharge of <b>\$130.00</b> for furnishing the oath or declaration later than 30 months from the earliest claimed priority date [37 CFR 1.492(e)]. |                     |               |                  |                 |
|   |   | Claims  | Number Filed        | Number Extra  | Rate             |                 |
|   |   | Total Claims  | 61 - 20 =           | 41            | <b>X \$18.00</b> | <b>\$738.00</b> |
|   |   | Independent Claims  | 11 - 3 =            | 8             | <b>X \$86.00</b> | <b>\$688.00</b> |
| Multiple dependent claim(s) (if applicable).  |   |   | + <b>\$290.00</b>   | <b>\$0.00</b> |                  |                 |
| <b>TOTAL OF ABOVE CALCULATIONS</b>  |   | <b>= \$1,526.00</b>   |                     |               |                  |                 |
| <input type="checkbox"/> Applicant claims small entity status. See 37 CFR 1.27. The fees indicated above are reduced by 1/2.  |   | <b>SUBTOTAL</b> = <b>\$1,526.00</b>   |                     |               |                  |                 |
| Processing fee of <b>\$130.00</b> for furnishing the English translation later than 30 months from the earliest claimed priority date [37 CFR 1.492(f)].  |   | +   |                     |               |                  |                 |
| <b>TOTAL NATIONAL FEE</b>   |   | <b>= \$1,526.00</b>   |                     |               |                  |                 |
| Fee for recording the enclosed assignment [37 CFR 1.21(h)]. The assignment must be accompanied by an appropriate cover sheet (37 CFR 3.28, 3.31). <b>\$40.00</b> per property.  |   | +   |                     |               |                  |                 |
| <b>TOTAL FEES ENCLOSED</b>  |   | <b>= \$1,526.00</b>   |                     |               |                  |                 |
|   |   | Amount to be refunded   |                     |               |                  |                 |
|   |   | charged   |                     |               |                  |                 |
| <p>a. <input type="checkbox"/> A check in the amount of \$ _____ to cover the above fees is enclosed.</p> <p>b. <input checked="" type="checkbox"/> Please charge my Deposit Account No. <u>13-2755</u> in the amount of <u>\$1,526.00</u> to cover the above fees. A duplicate copy of this sheet is enclosed.</p> <p>c. <input checked="" type="checkbox"/> The Commissioner is hereby authorized to charge any additional fees which may be required, or credit any overpayment to the Deposit Account No. <u>13-2755</u>. A duplicate copy of this sheet is enclosed.</p>   |   |   |                     |               |                  |                 |
| <p><b>NOTE: Where an appropriate time limit under 37 CFR 1.494 or 1.495 has not been met, a petition to revive [37 CFR 1.137(a) or (b)] must be filed and granted to restore the application to pending status.</b></p>   |   |   |                     |               |                  |                 |
| <p>SEND ALL CORRESPONDENCE TO:</p> <p><b>Customer No. 000210</b></p> <p>MERCK &amp; CO., INC.<br/>Patent Department, RY60-30<br/>P.O. Box 2000<br/>126 East Lincoln Avenue<br/>Rahway, New Jersey 07065-0970</p> <p><b>DATE: July 12, 2004</b></p> <p><b>PHONE #: (732) 594-0182</b></p>  |   |   |                     |               |                  |                 |
| <br><b>SIGNATURE</b><br><u>Raynard Yuro</u><br><b>NAME</b><br><b>45,570</b><br><b>REGISTRATION NUMBER</b>   |   |   |                     |               |                  |                 |